Customer No.: 000027683 Atty. Docket No.: 36697.17

## 1.2 AMENDMENTS TO THE CLAIMS (LISTING OF CLAIMS):

This listing of claims will replace all prior versions, and listings of claims in the application:

- 1. (Currently Amended) A method for assessing skeletal growth of a subject other than an adult in congestive heart failure, comprising measuring the level of NT-CNP in a biological samplefluid from the subject, and comparing the level against the this measured level of NT-CNP against a mean NT-CNP level from a control population for which at least a first skeletal growth information is known, wherein a significant deviation in the measured level of NT-CNP in the subject from the mean control level of NT-CNP in the control population is indicative of abnormal skeletal growth.
- 2. (Currently Amended) The method of claim 1, wherein the biological samplefluid is plasma or whole blood.
- 3. (Previously Presented) The method of claim 1, where said subject is a pre-adult.
- 4. (Previously Presented) The method of claim 1, wherein said subject is a pre-pubescent child or an infant.
- 5. (Currently Amended) The method of claim 3, wherein said subject is a neonate and the samplefluid comprises is a cord blood-sample.
- 6. (Previously Presented) The method of claim 1, wherein said subject is undergoing a treatment regimen, which may impact on skeletal growth in said subject.
- 7. (Previously Presented) The method of claim 1, wherein said subject is exposed to chemicals or other external factors which may impact on skeletal growth in said subject.

- 8. (Previously Presented) The method of claim 1, wherein said measuring step comprises detecting binding between NT-CNP and a binding agentan antibody that selectively binds NT-CNP.
- 9. (Currently Amended) The method of claim 8, wherein said binding agent is an antibody is an antibody fragment that selectively binds NT-CNP.
- 10. (Currently Amended) The method of claim <u>89</u>, wherein said <u>binding agentantibody</u> is a monoclonal antibody or <u>a monoclonal antibody</u> fragment.
- 11. (Currently Amended) The method of claim 8, wherein the NT-CNP to which the binding agentantibody selectively binds comprises an antigenic peptide selected from the group consisting of proCNP(1-103), proCNP(1-50), proCNP(1-81), and proCNP(51-81).
- 12. (Previously Presented) The method of claim 11, wherein the NT-CNP comprises proCNP(1-50).
- 13. (Currently Amended) The method of claim 8, wherein binding of NT-CNP is measured using with antibodies or antibody fragments that are immobilized to a solid phase.
- 14. (Currently Amended) A method for predicting the-skeletal growth potential of a subject other than an adult in congestive heart failure, comprising measuring the level of NT-CNP in a biological samplefluid from said subject, and comparing this measured level of the level against the mean NT-CNP against a mean NT-CNP level from the control population that has attained maximum skeletal growth and predicting from the NT-CNP level in the subject, skeletal growth potential of the subject.
- 15. (Currently Amended) A method for predicting the skeletal age of a subject other than an adult in congestive heart failure, comprising measuring the level of NT-CNP in a biological samplefluid from said subject and comparing this measured level of the level

against the mean NT-CNP against a mean NT-CNP level from level of a control population of known skeletal ages, and predicting from the NT-CNP level in the subject, the skeletal age of the subject.

- 16. (Currently Amended) A method for diagnosing a skeletal disease or disorder in a subject other than an adult in congestive heart failure, comprising measuring the level of NT-CNP in a biological samplefluid from said subject, and comparing this measured level of NT-CNP against a the level against the mean NT-CNP level from a control population, wherein a significant deviation in the measured level from the mean control level is indicative of a skeletal disease or disorder.
- 17. (Currently Amended) The method of claim 14, wherein said biological samplefluid is plasma or whole blood.
- 18. (Currently Amended) The method of claim 14, wherein said subject is a propre-adult.
- 19. (Previously Presented) The method of claim 14, wherein said subject is a pre-pubescent child or an infant.
- 20. (Currently Amended) The method of claim 16, wherein said subject is a neonate and said biological samplefluid comprises cord blood.
- 21. (Currently Amended) The method of claim 16, wherein the measuring step comprises detecting binding between NT-CNP and a binding agentan antibody or antibody fragment that selectively binds NT-CNP.
- 22. (Canceled)
- 23. (Currently Amended) The method of claim <u>2122</u>, wherein said <u>binding agentantibody or antibody fragment</u> is a monoclonal antibody or monoclonal antibody fragment.

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24. (Currently Amended) The method of claim 21, wherein the NT-CNP to which said binding agentantibody or antibody fragment selectively binds comprises an antigenic peptide selected from the group consisting of proCNP(1-103), proCNP(1-50), proCNP(1-81), and proCNP(51-81).

- 25. (Previously Presented) The method of claim 24, wherein said NT-CNP comprises proCNP(1-50).
- 26. (Currently Amended) The method of claim 21, wherein binding of said NT-CNP is measured usingwith an antibody or antibody fragment that isantibodies or antibody fragments that are immobilized to a solid phase.
- 27. (Currently Amended) The method of claim 26, wherein where a significant deviation from the mean control level is found in the samplefluid, the method comprises a further step of comparing the measured NT-CNP level with one or more mean NT-CNP levels from populations having known skeletal diseases or disorders to make a more accurate diagnosis of thea specific disease or disorder.
- 28. (Previously Presented) The method of claim 16, wherein said skeletal disease or disorder is selected from the group consisting of congenital disorders, delayed developmental disorders and advanced development syndromes.
- 29. (Currently Amended) A method of monitoring skeletal growth in a subject other than an adult in congestive heart failure, comprising:
  - (a) measuring the level of NT-CNP in a first biological samplefluid from said subject and measuring the level of NT-CNP in a second biological samplefluid, wherein said second biological samplefluid is taken from the same subject as said first samplebiological fluid but at a later date; and

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(b) comparing the levels of NT-CNP in said first and said second samplesbiological

fluids, wherein a significant change in the level of NT-CNP in said second

samplebiological fluid from the level of NT-CNP in said first samplebiological fluid

indicates a change in the rate of skeletal growth in said subject.

30. (Previously Presented) The method of claim 29, wherein said subject is undergoing a

treatment regimen that may impact skeletal growth of said subject.

31. (Currently Amended) The method of claim 6 or claim 30, wherein said treatment

regimen involves the administration of glucocorticoids to said subject.

32. (Previously Presented) The method of claim 31, wherein said subject is undergoing

treatment for asthma or other chronic allergic states.

33. (Currently Amended) A kit for assessing skeletal growth, diagnosing a skeletal disease or

disorder, or predicting skeletal growth potential or skeletal age in a subject other than an

adult in congestive heart failure, said kit comprising:

(a) means for measuring the level of NT-CNP in a biological sample fluid obtained from

said subject, comprising a binding agentan antibody or an antibody fragment that

selectively binds to a NT-CNP molecule selected from the group consisting of proCNP(1-

103), proCNP(1-50), proCNP(1-81), and proCNP(51-81), and which can be used to

quantitatively measure NT-CNP; and

(b) instructions for assessing or monitoring said skeletal growth, predicting said skeletal

growth potential or said skeletal age, or diagnosing said skeletal disease or disorder in

said subject from the NT-CNP level measured in said biological sample fluid.

34. (Canceled)

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35. (Currently Amended) The kit of claim [[34]]33, wherein said binding agentantibody or

said antibody fragment is a monoclonal antibody or a fragment thereof.

36. (Previously Presented) An NT-CNP binding agent that selectively binds a proCNP(1-

50), or proCNP(1750)a proCNP(1-103), a proCNP(1-50), a proCNP(1-81), or a

proCNP(51-81) peptide.

37-43. (Canceled)

Please add the following new claims:

44. (New) A method for assessing skeletal growth of a pre-adult subject, comprising

measuring NT-CNP in a biological fluid from the subject, and comparing this measured

level of NT-CNP against a mean NT-CNP level from a control population for which at

least a first skeletal growth information is known, wherein a significant deviation in the

measured level of NT-CNP in the subject from the mean level of NT-CNP in the control

population is indicative of abnormal skeletal growth.

45. (New) The method of claim 44, wherein said measuring step comprises detecting

binding between NT-CNP and an antibody or an antibody binding fragment that

selectively binds NT-CNP.

46. (New) The method of claim 45, wherein said antibody or said antibody binding fragment

selectively binds an NT-proCNP peptide.

47. (New) The method of claim 46, wherein said NT-proCNP peptide comprises an

antigenic peptide selected from the group consisting of proCNP(1-103), proCNP(1-50),

proCNP(1-81), and proCNP(51-81).

- 48. (New) A method for assessing skeletal growth of a subject suspected of having a skeletal disease or disorder, comprising measuring NT-CNP in a biological fluid from the subject, and comparing this measured level of NT-CNP against a mean NT-CNP level from a control population for which at least a first skeletal growth information is known, wherein a significant deviation in the measured level of NT-CNP in the subject from the mean level of NT-CNP in the control population is indicative of abnormal skeletal growth in said subject.
- 49. (New) The method of claim 48, wherein said measuring step comprises detecting binding between NT-CNP and an antibody or an antibody binding fragment that selectively binds NT-CNP.
- 50. (New) The method of claim 49, wherein said antibody or said antibody binding fragment selectively binds an NT-proCNP peptide.
- 51. (New) The method of claim 50, wherein said NT-proCNP peptide comprises an antigenic peptide selected from the group consisting of proCNP(1-103), proCNP(1-50), proCNP(1-81), and proCNP(51-81).